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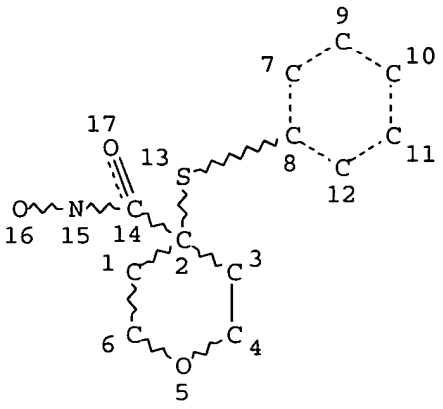

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IN **2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4'-(4-pyridinyl)[1,1'-biphenyl]-4-yl]sulfonyl]- (9CI)**
MF C23 H22 N2 O5 S
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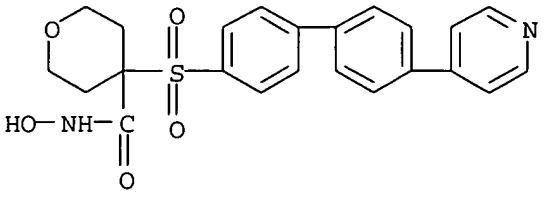

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FILE 'CAPLUS' ENTERED AT 12:28:54 ON 18 APR 2005
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FILE COVERS 1907 - 18 Apr 2005 VOL 142 ISS 17
FILE LAST UPDATED: 17 Apr 2005 (20050417/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

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TI   Preparation of arylsulfonylheterocyclylhydroxamic acids and related
     compounds as matrix metalloprotease inhibitors
IN   Barta, Thomas E.; Becker, Daniel P.; Boehm, Terri L.; De Crescenzo, Gary
     A.; Villamil, Clara I.; McDonald, Joseph J.; Freskos, John N.; Getman,
     Daniel P.
PA   G.D. Searle and Co., USA
SO   PCT Int. Appl., 840 pp.
     CODEN: PIXXD2
DT   Patent
LA   English
FAN.CNT 5
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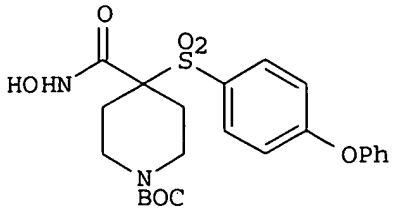

I

AB A process for treating conditions associated with pathol. matrix metalloproteinase (MMP) activity comprises administration of compds. having inhibitory activity against >1 of MMP-2, MMP-9, and MMP-13, while exhibiting substantially less inhibition of MMP-1. The compds. are of the form HONHCOCR1R2SO2R3 [R1, R2 = H; R1R2 = atoms to form a 5-8 membered ring containing 1-3 heteroatoms; R3 = (substituted) aryl, heteroaryl]. Thus, 4-PhOC6H4SH was heated in Me2SO to give the disulfide dimer, which in THF was added to a mixture of Et N-tert-butoxycarbonylisonipecotate (preparation given) and LDA in THF at -60° to room temperature to give 405 sulfide, which was oxidized with m-ClC6H4CO(OOH) to give 59% sulfone. The Et ester was saponified with NaOH in EtOH/H2O to give 100% acid, which in DMF was treated with hydroxybenzotriazole, EDC, 4-methylmorpholine, and aqueous NH2OH to give title compound (I). I inhibited MMP-2 with IC50 = 0.2 nM.

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(preparation of arylsulfonylheterocyclylhydroxamic acids and related compds. as matrix metalloprotease inhibitors)

RN 226390-38-1 CAPLUS

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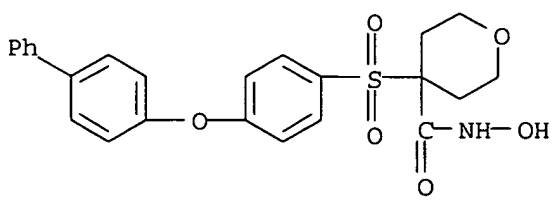

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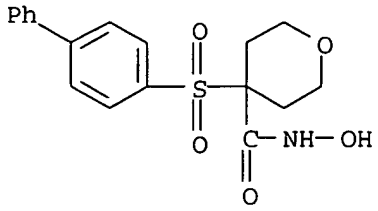

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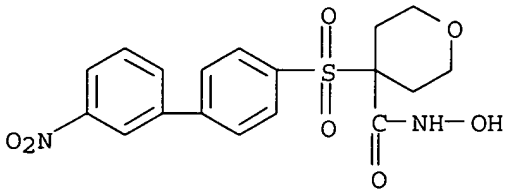

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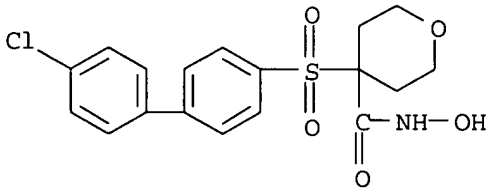

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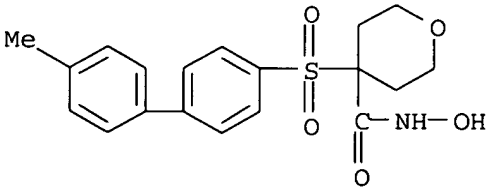

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CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4'-[2-(1-pyrrolidinyl)ethoxy][1,1'-biphenyl]-4-yl]sulfonyl]-, mono(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

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CRN 226393-42-6
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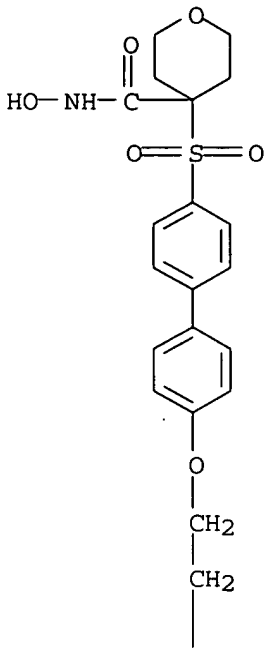

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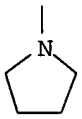

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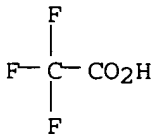

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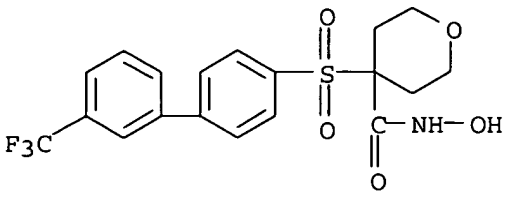

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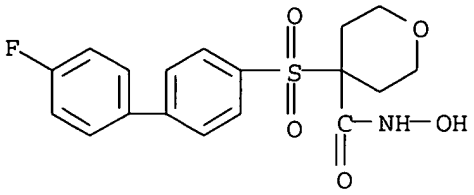

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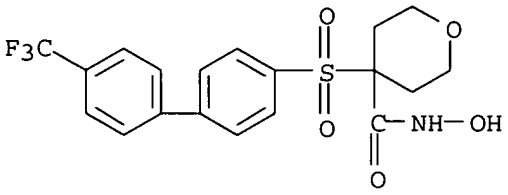

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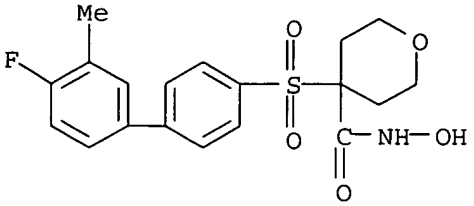

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CN 2H-Pyran-4-carboxamide, 4-[(3',4'-dimethyl[1,1'-biphenyl]-4-yl)sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

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CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[3'-(propylamino)[1,1'-biphenyl]-4-yl]sulfonyl]-, mono(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

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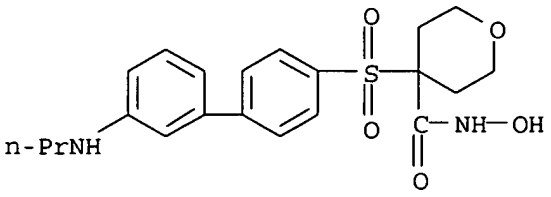

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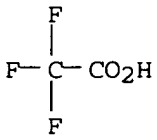

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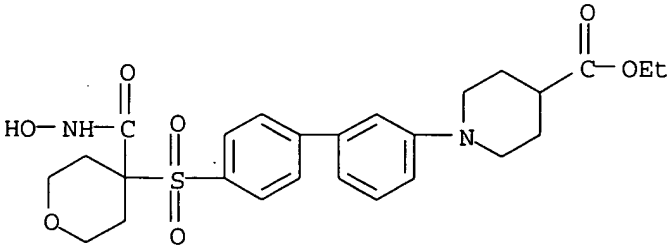

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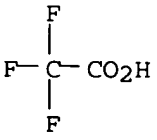

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L13 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN
AN 2000:608722 CAPLUS
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IN Barta, Thomas E.; Becker, Daniel P.; Bedell, Louis J.; Boehm, Terri L.; Carroll, Jeffery N.; De Crescenzo, Gary A.; Fobian, Yvette M.; Freskos, John N.; Getman, Daniel P.; McDonald, Joseph J.; Hanson, Gunnar J.; Hockerman, Susan L.; Howard, Susan C.; Kolodziej, Steve A.; Li, Hui; Mischke, Deborah A.; Rico, Joseph G.; Stehle, Nathan W.; Tollefson, Michael B.; Vernier, William F.; Villamil, Clara I.; Rao, Shashidahar N.
PA G.D. Searle and Co., USA
SO PCT Int. Appl., 851 pp.
CODEN: PIXXD2
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(preparation of arylsulfonylheterocyclylhydroxamic acids and related compds. as matrix metalloprotease inhibitors)

RN 226390-38-1 CAPLUS
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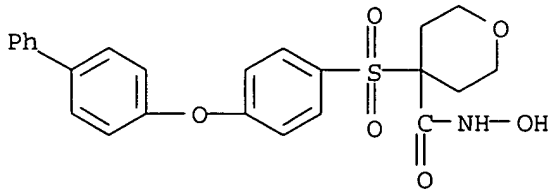

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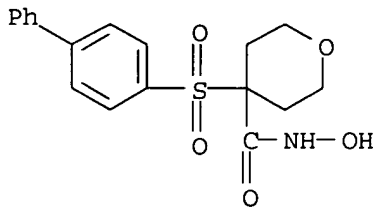

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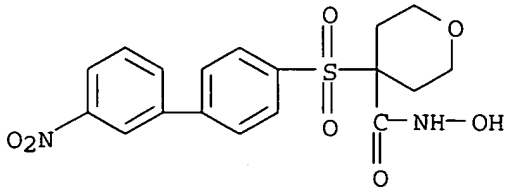

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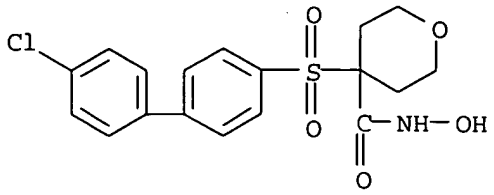

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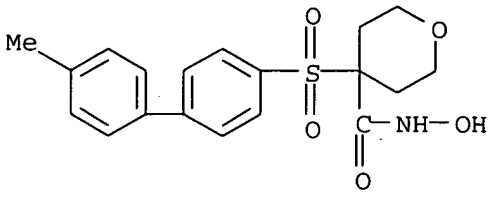
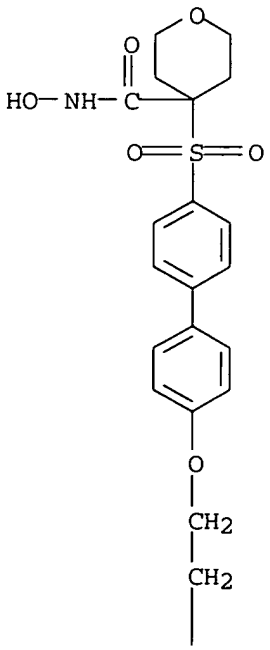


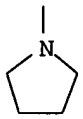
RN 226393-43-7 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4'-[2-(1-pyrrolidinyl)ethoxy][1,1'-biphenyl]-4-yl]sulfonyl]-, mono(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 226393-42-6
CMF C24 H30 N2 O6 S

PAGE 1-A





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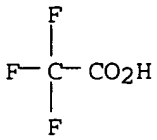

RN 226394-47-4 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[3'-(1-piperazinyl)[1,1'-biphenyl]-4-yl]sulfonyl]-, bis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

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CRN 226394-46-3
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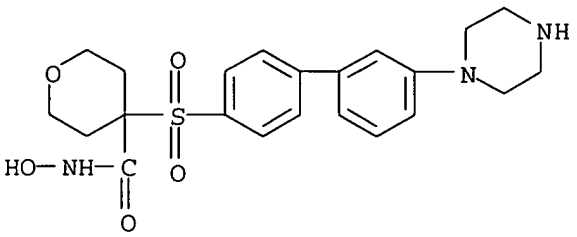

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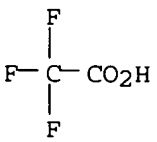

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L13 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN
AN 2000:824220 CAPLUS
DN 134:17399
TI Aromatic sulfone hydroxamic acid metalloprotease inhibitors
IN Barta, Thomas E.; Becker, Daniel P.; Bedell, Louis J.; Boehm, Terri L.; Carroll, Jeffrey N.; Decrescenzo, Gary A.; Fobian, Yvette M.; Freskos, John N.; Getman, Daniel P.; McDonald, Joseph J.; Hockerman, Susan L.; Howard, Susan C.; Kolodziej, Stephen A.; Li, Madeleine Hui; Mischke, Deborah A.; Rico, Joseph G.; Stehle, Nathan W.; Tollefson, Michael B.; Vernier, William F.; Villamil, Clara I.
PA G.D. Searle and Co., USA
SO PCT Int. Appl., 616 pp.

CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 5

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2000069821	A1	20001123	WO 2000-US6719	20000515 <--
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RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6750228	B1	20040615	US 2000-570731	20000512
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EP 1183239	A1	20020306	EP 2000-930088	20000515
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BR 2000010562	A	20030610	BR 2000-10562	20000515
JP 2003520196	T2	20030702	JP 2000-618238	20000515
AU 766792	B2	20031023	AU 2000-47970	20000515
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ZA 2001009006	A	20021202	ZA 2001-9006	20011031
NO 2001005543	A	20020110	NO 2001-5543	20011113
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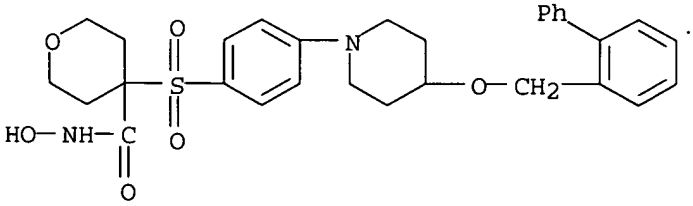
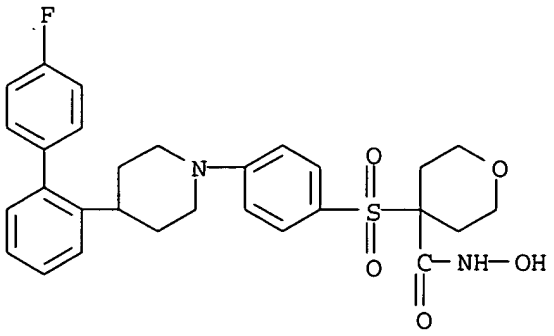
US 2000-570731	A	20000512		
US 1997-66007P	P	19971114		
US 1998-95347P	P	19980804		
US 1998-101080P	P	19980918		
US 1999-256948	B2	19990224		
WO 2000-US6719	W	20000515		

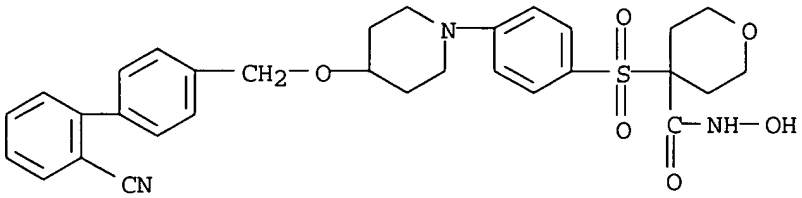
OS MARPAT 134:17399
IT **308823-70-3P 308827-37-4P 308827-51-2P**
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(drug candidate; preparation of aromatic sulfone hydroxamic acids as metalloprotease inhibitors)
RN 308823-70-3 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[4-([1,1'-biphenyl]-2-ylmethoxy)-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



RN 308827-37-4 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[4-(4'-fluoro[1,1'-biphenyl]-2-yl)-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



RN 308827-51-2 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[4-[(2'-cyano[1,1'-biphenyl]-4-yl)methoxy]-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L13 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN
AN 2004:485162 CAPLUS
DN 141:38534
TI Preparation of aromatic sulfone hydroxamic acid metalloprotease inhibitors
IN Barta, Thomas E.; Becker, Daniel P.; Bedell, Louis J.; Boehm, Terri L.; Carroll, Jeffrey N.; Decrescenzo, Gary A.; Fobian, Yvette M.; Freskos, John N.; Getman, Daniel P.; McDonald, Joseph J.; Li, Madeleine H.; Hockerman, Susan L.; Howard, Susan C.; Kolodziej, Steve A.; Mischke, Deborah A.; Rico, Joseph G.; Stehle, Nathan W.; Tollefson, Michael B.; Vernier, William F.; Villamil, Clara I.
PA Pharmacia Corporation, USA
SO U.S., 403 pp., Cont.-in-part of U.S. Ser. No. 311,837.
CODEN: USXXAM
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LA English
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	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
PI	US 6750228	B1	20040615	US 2000-570731	20000512	
	US 2001014688	A1	20010816	US 1998-191129	19981113	<--
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EP 1183239	A1	20020306	EP 2000-930088		20000515
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US 2003073718	A1	20030417	US 2001-989943		20011121
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US 2000-570731	A	20000512			
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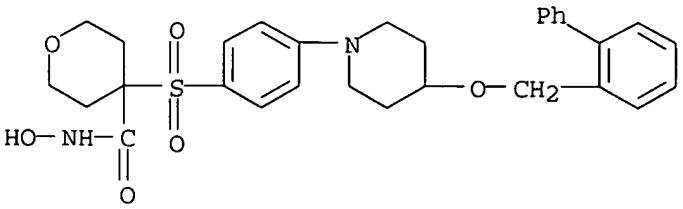
OS MARPAT 141:38534

IT **308823-70-3P 308827-37-4P 308827-51-2P**
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(drug candidate; preparation of aromatic sulfone hydroxamic acids as metalloprotease inhibitors)

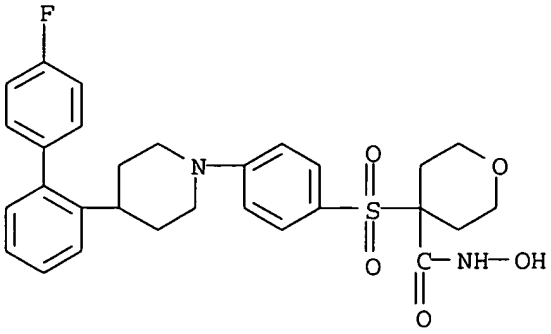
RN 308823-70-3 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[4-([1,1'-biphenyl]-2-ylmethoxy)-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

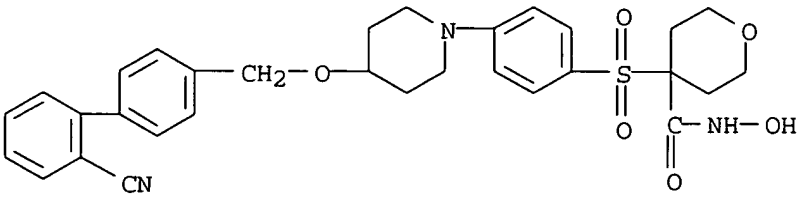


RN 308827-37-4 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[4-(4'-fluoro[1,1'-biphenyl]-2-yl)-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



RN 308827-51-2 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[4-[(2'-cyano[1,1'-biphenyl]-4-yl)methoxy]-1-piperidinyl]phenyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)



RE.CNT 54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

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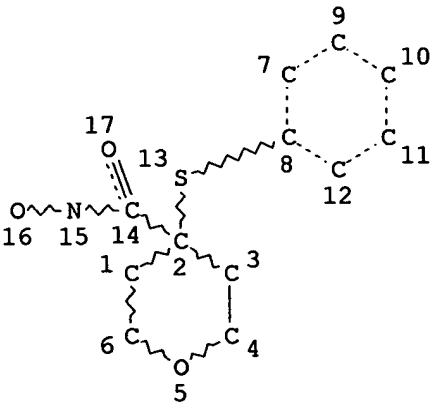

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STEREO ATTRIBUTES: NONE

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